

## SEQUENCE LISTING

<110> RYBAK, SUSANNA M. GOLDENBERG, DAVID M. NEWTON, DIANNE L.

<120> IMMUNOCONJUGATES OF TOXINS DIRECTED AGAINST MALIGNANT CELLS

<130> 018733/1059

<140> 09/961,400

<141> 2001-09-25

<150> 09/622,613

<151> 2000-08-17

<150> PCT/US99/06641

<151> 1999-03-26

<150> 60/079,751

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<170> PatentIn Ver. 2.1

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gtt gac tgt aat aat atc atg tca aca aac ttg ttc cac tgc aag gac 96
Val Asp Cys Asn Asn Ile Met Ser Thr Asn Leu Phe His Cys Lys Asp
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aag aac act ttt atc tat tca cgt cct gag cca gtg aag gcc atc tgt 144 Lys Asn Thr Phe Ile Tyr Ser Arg Pro Glu Pro Val Lys Ala Ile Cys 35 40 45

aaa gga att ata gcc tcc aaa aat gtg tta act acc tct gag ttt tat 192 Lys Gly Ile Ile Ala Ser Lys Asn Val Leu Thr Thr Ser Glu Phe Tyr 50 55 60

ctc tct gat tgc aat gta aca agc agg cct tgc aag tat aaa tta aag 240 Leu Ser Asp Cys Asn Val Thr Ser Arg Pro Cys Lys Tyr Lys Leu Lys 65 70 75 80

## BEST AVAILABLE COPY

2

aaa tca act aat aca ttt tgt gta act tgt gag aat caa gct cca gta Lys Ser Thr Asn Thr Phe Cys Val Thr Cys Glur Asn Gln Ala Pro Val 85 90 95 cat ttc gtg ggt gtc gga cat tgc 312 His Phe Val Gly Val Gly His Cys 100 <210> 2 <211> 104 <212> PRT <213> Rana pipiens <400> 2 Gln Asp Trp Leu Thr Phe Gln Lys Lys His Leu Thr Asn Thr Arg Asp Val Asp Cys Asn Asn Ile Met Ser Thr Asn Leu Phe His Cys Lys Asp Lys Asn Thr Phe Ile Tyr Ser Arg Pro Glu Pro Val Lys Ala Ile Cys 40 35 Lys Gly Ile Ile Ala Ser Lys Asn Val Leu Thr Thr Ser Glu Phe Tyr Leu Ser Asp Cys Asn Val Thr Ser Arg Pro Cys Lys Tyr Lys Leu Lys Lys Ser Thr Asn Thr Phe Cys Val Thr Cys Glu Asn Gln Ala Pro Val 90 85 His Phe Val Gly Val Gly His Cys 100 <210> 3 <211> 312 <212> DNA <213> Rana pipiens <220> <221> CDS <222> (1)..(312) <400> 3 caa gac tgg ctt acg ttt cag aag aag cac ctg aca aac acc cgg gat Gln Asp Trp Leu Thr Phe Gln Lys Lys His Leu Thr Asn Thr Arg Asp 10 gtt gac tgt aat aat atc ctg tca aca aac ttg ttc cac tgc aag gac Val Asp Cys Asn Asn Ile Leu Ser Thr Asn Leu Phe His Cys Lys Asp

25

B)

**√**17

aag aac act ttt atc tat tca cgt cct gag cca gtg aag gcc atc tgt Lys Asn Thr Phe Ile Tyr Ser Arg Pro Glu Pro Val Lys Ala Ile Cys 40 aaa gga att ata gcc tcc aaa aat gtg tta act acc ttt gag ttt tat 192 Lys Gly Ile Ile Ala Ser Lys Asn Val Leu Thr Thr Phe Glu Phe Tyr 55 ctc tct gat tgc aat gta aca agc agg cct tgc aag tat aaa tta aag Leu Ser Asp Cys Asn Val Thr Ser Arg Pro Cys Lys Tyr Lys Leu Lys 70 aaa tca act aat aca ttt tgt gta act tgt gag aat caa gct cca gta Lys Ser Thr Asn Thr Phe Cys Val Thr Cys Glu Asn Gln Ala Pro Val 312 cat ttc gtg ggt gtc gga cat tgc His Phe Val Gly Val Gly His Cys <210> 4 <211> 104 <212> PRT <213> Rana pipiens <400> 4 Gln Asp Trp Leu Thr Phe Gln Lys Lys His Leu Thr Asn Thr Arg Asp Val Asp Cys Asn Asn Ile Leu Ser Thr Asn Leu Phe His Cys Lys Asp 25 Lys Asn Thr Phe Ile Tyr Ser Arg Pro Glu Pro Val Lys Ala Ile Cys

B1 cut

Lys Gly Ile Ile Ala Ser Lys Asn Val Leu Thr Thr Phe Glu Phe Tyr 50 60

Leu Ser Asp Cys Asn Val Thr Ser Arg Pro Cys Lys Tyr Lys Leu Lys 65 70 75 80

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His Phe Val Gly Val Gly His Cys

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gat gtt gac tgt aat aat atc ctg tca aca aac ttg ttc cac tgc aag
                                                                   96
Asp Val Asp Cys Asn Asn Ile Leu Ser Thr Asn Leu Phe His Cys Lys
gac aag aac act ttt atc tat tca cgt cct gag cca gtg aag gcc atc
Asp Lys Asn Thr Phe Ile Tyr Ser Arg Pro Glu Pro Val Lys Ala Ile
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tgt aaa gga att ata gcc tcc aaa aat gtg tta act acc ttt gag ttt
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Cys Lys Gly Ile Ile Ala Ser Lys Asn Val Leu Thr Thr Phe Glu Phe
     50
tat ctc tct gat tgc aat gca aca agc agg cct tgc aag tat aaa tta
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Tyr Leu Ser Asp Cys Asn Ala Thr Ser Arg Pro Cys Lys Tyr Lys Leu
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                     70
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Lys Lys Ser Thr Asn Thr Phe Cys Val Thr Cys Glu Asn Gln Ala Pro
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gta cat ttc gtg ggt gtc gga cat tgc
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Cys Lys Gly Ile Ile Ala Ser Lys Asn Val Leu Thr Thr Phe Glu Phe
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Tyr Leu Ser Asp Cys Asn Ala Thr Ser Arg Pro Cys Lys Tyr Lys Leu
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Asn Leu Phe His Cys Lys Asp Lys Asn Thr Phe Ile Tyr Ser Arg Pro
Glu Pro Val Lys Ala Ile Cys Lys Gly Ile Ile Ala Ser Lys Asn Val
Leu Thr Thr Ser Glu Phe Tyr Leu Ser Asp Cys Asn Val Thr Ser Arg
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gtt gac tgt aat aat atc atg tca aca aac ttg ttc cac tgc aag gac Val Asp Cys Asn Asn Ile Met Ser Thr Asn Leu Phe His Cys Lys Asp

10

aag aac act ttt atc tat tca cgt cct gag cca gtg aag gcc atc tgt Lys Asn Thr Phe Ile Tyr Ser Arg Pro Glu Pro Val Lys Ala Ile Cys 40 aaa gga att ata gcc tcc aaa aat gtg tta act acc tct gag ttt tat 192 Lys Gly Ile Ile Ala Ser Lys Asn Val Leu Thr Thr Ser Glu Phe Tyr 55 ctc tct gat tgc aat gta aca agc agg cct tgc aag tat aaa tta aag Leu Ser Asp Cys Asn Val Thr Ser Arg Pro Cys Lys Tyr Lys Leu Lys aaa tca act aat aca ttt tgt gta act tgt gag aat caa gct cca gta 288 Lys Ser Thr Asn Thr Phe Cys Val Thr Cys Glu Asn Gln Ala Pro Val 85 312 cat ttc gtg ggt gtc gga cat tgc His Phe Val Gly Val Gly His Cys <210> 11 <211> 104 <212> PRT <213> Rana pipiens <400> 11 Ser Asp Trp Leu Thr Phe Gln Lys Lys His Leu Thr Asn Thr Arg Asp Val Asp Cys Asn Asn Ile Met Ser Thr Asn Leu Phe His Cys Lys Asp 25 Lys Asn Thr Phe Ile Tyr Ser Arg Pro Glu Pro Val Lys Ala Ile Cys Lys Gly Ile Ile Ala Ser Lys Asn Val Leu Thr Thr Ser Glu Phe Tyr Leu Ser Asp Cys Asn Val Thr Ser Arg Pro Cys Lys Tyr Lys Leu Lys Lys Ser Thr Asn Thr Phe Cys Val Thr Cys Glu Asn Gln Ala Pro Val 90 His Phe Val Gly Val Gly His Cys 100 <210> 12 <211> 315 <212> DNA <213> Rana pipiens <220> <221> CDS <222> (1)..(315)

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| atg tca gac<br>Met Ser Asp<br>1  |  |                                  |                          |                                |                                |                                |                                |                         |                         |                                |                    | 48  |
| gat gtt gac<br>Asp Val Asp   |  |                                  |                          |                                |                                |                                |                                |                         |                         |                                |                    | 96  |
| gac aag aac<br>Asp Lys Asn<br>35   |  |                                  |                          |                                |                                |                                |                                |                         |                         |                                |                    | 144 |
| tgt aaa gga<br>Cys Lys Gly<br>50   |  |                                  |                          |                                |                                |                                |                                |                         |                         |                                |                    | 192 |
| tat ctc tct<br>Tyr Leu Ser<br>65   |  |                                  |                          |                                |                                |                                |                                |                         |                         |                                |                    | 240 |
| aag aaa tca<br>Lys Lys Ser   |  |                                  |                          |                                |                                |                                |                                |                         |                         |                                |                    | 288 |
| gta cat ttc<br>Val His Phe   |  |                                  |                          |                                |                                |                                |                                |                         |                         |                                |                    | 315 |
|  |  |                                  |                          |                                |                                |                                |                                |                         |                         |                                |                    |     |
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| <211> 105<br><212> PRT<br><213> Rana p<br><400> 13<br>Met Ser Asp                        | Trp Leu                                      |                                  |                          | Lys                            | 10                             |                                |                                |                         |                         | 15                             |                    |     |
| <211> 105<br><212> PRT<br><213> Rana p<br><400> 13<br>Met Ser Asp<br>1                   | Trp Leu<br>5<br>Cys Asn<br>20                | Asn I                            | le Met                   | Lys<br>Ser<br>25               | 10<br>Thr                      | Asn                            | Leu                            | Phe                     | His<br>30               | 15<br>Cys                      | Lys                |     |
| <pre>&lt;211&gt; 105 &lt;212&gt; PRT &lt;213&gt; Rana p &lt;400&gt; 13 Met Ser Asp</pre> | Trp Leu<br>5<br>Cys Asn<br>20<br>Thr Phe     | Asn I                            | le Met<br>Cyr Ser<br>40  | Lys<br>Ser<br>25<br>Arg        | 10<br>Thr<br>Pro               | Asn<br>Glu                     | Leu<br>Pro                     | Phe<br>Val<br>45        | His<br>30<br>Lys        | 15<br>Cys<br>Ala               | Lys<br>Ile         |     |
| <pre>&lt;211&gt; 105 &lt;212&gt; PRT &lt;213&gt; Rana p &lt;400&gt; 13 Met Ser Asp</pre> | Trp Leu 5 Cys Asn 20 Thr Phe                 | Asn I                            | Tyr Ser<br>40<br>Ger Lys | Lys<br>Ser<br>25<br>Arg<br>Asn | 10<br>Thr<br>Pro<br>Val        | Asn<br>Glu<br>Leu              | Leu<br>Pro<br>Thr              | Phe<br>Val<br>45<br>Thr | His<br>30<br>Lys<br>Ser | 15<br>Cys<br>Ala<br>Glu        | Lys<br>Ile<br>Phe  |     |
| <pre>&lt;211&gt; 105 &lt;212&gt; PRT &lt;213&gt; Rana p &lt;400&gt; 13 Met Ser Asp</pre> | Trp Leu 5 Cys Asn 20 Thr Phe Ile Ile Asp Cys | Asn I Ile T Ala S Asn V 70 Thr P | Thr                      | Lys<br>Ser<br>25<br>Arg<br>Asn | 10<br>Thr<br>Pro<br>Val<br>Arg | Asn<br>Glu<br>Leu<br>Pro<br>75 | Leu<br>Pro<br>Thr<br>60<br>Cys | Phe Val 45 Thr          | His<br>30<br>Lys<br>Ser | 15<br>Cys<br>Ala<br>Glu<br>Lys | Lys Ile Phe Leu 80 |     |

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atc tgc aac act atc atg gac aac atc tac atc gtt ggt ggt cag
Ile Cys Asn Thr Ile Met Asp Asn Asn Ile Tyr Ile Val Gly Gln
tgc aaa cgt gtt aac act ttc atc atc tct tct gct act act gtt aaa
Cys Lys Arg Val Asn Thr Phe Ile Ile Ser Ser Ala Thr Thr Val Lys
         35
gct atc tgc act ggt gtt atc aac atg aac gtt ctg tct act act cgt
                                                                  192
Ala Ile Cys Thr Gly Val Ile Asn Met Asn Val Leu Ser Thr Thr Arg
     50
                         55
ttc cag ctg aac act tgc act cgt act tct atc act ccg cgt ccg tgc
Phe Gln Leu Asn Thr Cys Thr Arg Thr Ser Ile Thr Pro Arg Pro Cys
ccg tac tct tct cgt act gaa act aac tac atc tgc gtt aaa tgc gaa
Pro Tyr Ser Ser Arg Thr Glu Thr Asn Tyr Ile Cys Val Lys Cys Glu
aac cag tac ccg gtt cat ttc gct ggt atc ggt cgt tgc ccg
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Asn Gln Tyr Pro Val His Phe Ala Gly Ile Gly Arg Cys Pro
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Ile Cys Asn Thr Ile Met Asp Asn Asn Ile Tyr Ile Val Gly Gln
Cys Lys Arg Val Asn Thr Phe Ile Ile Ser Ser Ala Thr Thr Val Lys
Ala Ile Cys Thr Gly Val Ile Asn Met Asn Val Leu Ser Thr Thr Arg
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BI

Phe Gln Leu Asn Thr Cys Thr Arg Thr Ser Ile Thr Pro Arg Pro Cys

Pro Tyr Ser Ser Arg Thr Glu Thr Asn Tyr Ile Cys Val Lys Cys Glu Asn Gln Tyr Pro Val His Phe Ala Gly Ile Gly Arg Cys Pro <210> 16 <211> 333 <212> DNA <213> Rana catesbeiana <220> <221> CDS <222> (1)..(333) <400> 16 atg cag aac tgg gct act ttc cag cag aaa cat atc atc aac act ccg Met Gln Asn Trp Ala Thr Phe Gln Gln Lys His Ile Ile Asn Thr Pro 5 atc atc tgc aac act atc atg gac aac atc tac atc gtt ggt ggt 96 Ile Ile Cys Asn Thr Ile Met Asp Asn Asn Ile Tyr Ile Val Gly Gly cag tgc aaa cgt gtt aac act ttc atc atc tct tct gct act act gtt 144 Gln Cys Lys Arg Val Asn Thr Phe Ile Ile Ser Ser Ala Thr Thr Val 35 40 aaa gct atc tgc act ggt gtt atc aac atg aac gtt ctg tct act act 192 Lys Ala Ile Cys Thr Gly Val Ile Asn Met Asn Val Leu Ser Thr Thr cqt ttc cag ctg aac act tgc act cgt act tct atc act ccg cgt ccg Arg Phe Gln Leu Asn Thr Cys Thr Arg Thr Ser Ile Thr Pro Arg Pro 70 tgc ccg tac tct tct cgt act gaa act aac tac atc tgc gtt aaa tgc 288 Cys Pro Tyr Ser Ser Arg Thr Glu Thr Asn Tyr Ile Cys Val Lys Cys gaa aac cag tac ccg gtt cat ttc gct ggt atc ggt cgt tgc ccg 333 Glu Asn Gln Tyr Pro Val His Phe Ala Gly Ile Gly Arg Cys Pro 105 <210> 17 <211> 111 <212> PRT <213> Rana catesbeiana

Bij

Met Gln Asn Trp Ala Thr Phe Gln Gln Lys His Ile Ile Asn Thr Pro

Ile Ile Cys Asn Thr Ile Met Asp Asn Asn Ile Tyr Ile Val Gly Gly Gln Cys Lys Arg Val Asn Thr Phe Ile Ile Ser Ser Ala Thr Thr Val Lys Ala Ile Cys Thr Gly Val Ile Asn Met Asn Val Leu Ser Thr Thr 55 Arg Phe Gln Leu Asn Thr Cys Thr Arg Thr Ser Ile Thr Pro Arg Pro Cys Pro Tyr Ser Ser Arg Thr Glu Thr Asn Tyr Ile Cys Val Lys Cys Glu Asn Gln Tyr Pro Val His Phe Ala Gly Ile Gly Arg Cys Pro 105 100 <210> 18 <211> 330 <212> DNA <213> Rana catesbeiana <220> <221> CDS <222> (1)..(330) <400> 18 cag aac tgg gct act ttc cag cag aaa cat atc atc aac act ccg atc 48 Gln Asn Trp Ala Thr Phe Gln Gln Lys His Ile Ile Asn Thr Pro Ile atc tgc aac act atc ctg gac aac atc tac atc gtt ggt ggt cag Ile Cys Asn Thr Ile Leu Asp Asn Asn Ile Tyr Ile Val Gly Gln 25 20 tgc aaa cgt gtt aac act ttc atc tct tct gct act act gtt aaa Cys Lys Arg Val Asn Thr Phe Ile Ile Ser Ser Ala Thr Thr Val Lys 40 get ate tge act ggt gtt ate aac etg aac gtt etg tet act act egt 192 Ala Ile Cys Thr Gly Val Ile Asn Leu Asn Val Leu Ser Thr Thr Arg ttc cag ctg aac act tgc act cgt act tct atc act ccg cgt ccg tgc 240 Phe Gln Leu Asn Thr Cys Thr Arg Thr Ser Ile Thr Pro Arg Pro Cys ceg tac tet tet egt act gaa act aac tac ate tge gtt aaa tge gaa Pro Tyr Ser Ser Arg Thr Glu Thr Asn Tyr Ile Cys Val Lys Cys Glu

85

By

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Ile Cys Asn Thr Ile Leu Asp Asn Asn Ile Tyr Ile Val Gly Gln
Cys Lys Arg Val Asn Thr Phe Ile Ile Ser Ser Ala Thr Thr Val Lys
Ala Ile Cys Thr Gly Val Ile Asn Leu Asn Val Leu Ser Thr Thr Arg
                         55
Phe Gln Leu Asn Thr Cys Thr Arg Thr Ser Ile Thr Pro Arg Pro Cys
Pro Tyr Ser Ser Arg Thr Glu Thr Asn Tyr Ile Cys Val Lys Cys Glu
Asn Gln Tyr Pro Val His Phe Ala Gly Ile Gly Arg Cys Pro
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ate ate tge aac act ate etg gae aac ate tae ate gtt ggt ggt
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Ile Ile Cys Asn Thr Ile Leu Asp Asn Asn Ile Tyr Ile Val Gly Gly
20 25 30

cag tgc aaa cgt gtt aac act ttc atc atc tct tct gct act act gtt Gln Cys Lys Arg Val Asn Thr Phe Ile Ile Ser Ser Ala Thr Thr Val

aaa gct atc tgc act ggt gtt atc aac ctg aac gtt ctg tct act act Lys Ala Ile Cys Thr Gly Val Ile Asn Leu Asn Val Leu Ser Thr Thr 55 cgt ttc cag ctg aac act tgc act cgt act tct atc act ccg cgt ccg Arg Phe Gln Leu Asn Thr Cys Thr Arg Thr Ser Ile Thr Pro Arg Pro tgc ccg tac tct tct cgt act gaa act aac tac atc tgc gtt aaa tgc 288 Cys Pro Tyr Ser Ser Arg Thr Glu Thr Asn Tyr Ile Cys Val Lys Cys 90 333 gaa aac cag tac ccg gtt cat ttc gct ggt atc ggt cgt tgc ccg Glu Asn Gln Tyr Pro Val His Phe Ala Gly Ile Gly Arg Cys Pro <210> 21 <211> 111 <212> PRT <213> Rana catesbeiana <400> 21 Met Gln Asn Trp Ala Thr Phe Gln Gln Lys His Ile Ile Asn Thr Pro Ile Ile Cys Asn Thr Ile Leu Asp Asn Asn Ile Tyr Ile Val Gly Gly 25 Gln Cys Lys Arg Val Asn Thr Phe Ile Ile Ser Ser Ala Thr Thr Val 40 Lys Ala Ile Cys Thr Gly Val Ile Asn Leu Asn Val Leu Ser Thr Thr 55 Arg Phe Gln Leu Asn Thr Cys Thr Arg Thr Ser Ile Thr Pro Arg Pro Cys Pro Tyr Ser Ser Arg Thr Glu Thr Asn Tyr Ile Cys Val Lys Cys Glu Asn Gln Tyr Pro Val His Phe Ala Gly Ile Gly Arg Cys Pro 105 100 <210> 22 <211> 117 <212> PRT <213> Rana catesbeiana His His His His His Met Gln Asn Trp Ala Thr Phe Gln Gln Lys

His Ile Ile Asn Thr Pro Ile Ile Cys Asn Thr Ile Leu Asp Asn Asn 20 25 30

BI

Ile Tyr Ile Val Gly Gly Gln Cys Lys Arg Val Asn Thr Phe Ile Ile 40 Ser Ser Ala Thr Thr Val Lys Ala Ile Cys Thr Gly Val Ile Asn Leu Asn Val Leu Ser Thr Thr Arg Phe Gln Leu Asn Thr Cys Thr Arg Thr Ser Ile Thr Pro Arg Pro Cys Pro Tyr Ser Ser Arg Thr Glu Thr Asn Tyr Ile Cys Val Lys Cys Glu Asn Gln Tyr Pro Val His Phe Ala Gly 100 105 Ile Gly Arg Cys Pro 115 <210> 23 <211> 330 <212> DNA <213> Rana catesbeiana <220> <221> CDS <222> (1) .. (330) <400> 23 tca aac tgg gct act ttc cag cag aaa cat atc atc aac act ccg atc Ser Asn Trp Ala Thr Phe Gln Gln Lys His Ile Ile Asn Thr Pro Ile 10 atc tgc aac act atc atg gac aac atc tac atc gtt ggt ggt cag Ile Cys Asn Thr Ile Met Asp Asn Ile Tyr Ile Val Gly Gln tgc aaa cgt gtt aac act ttc atc atc tct tct gct act act gtt aaa 144 Cys Lys Arg Val Asn Thr Phe Ile Ile Ser Ser Ala Thr Thr Val Lys get ate tge act ggt gtt ate aac atg aac gtt etg tet act act egt Ala Ile Cys Thr Gly Val Ile Asn Met Asn Val Leu Ser Thr Thr Arg 50 tte caq etq aac act tge act egt act tet atc act eeg egt eeg tge Phe Gln Leu Asn Thr Cys Thr Arg Thr Ser Ile Thr Pro Arg Pro Cys 70 65 ccg tac tct tct cgt act gaa act aac tac atc tgc gtt aaa tgc gaa Pro Tyr Ser Ser Arg Thr Glu Thr Asn Tyr Ile Cys Val Lys Cys Glu 85 90 aac cag tac ccg gtt cat ttc gct ggt atc ggt cgt tgc ccg 330 Asn Gln Tyr Pro Val His Phe Ala Gly Ile Gly Arg Cys Pro

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Ile Cys Asn Thr Ile Met Asp Asn Asn Ile Tyr Ile Val Gly Gln
Cys Lys Arg Val Asn Thr Phe Ile Ile Ser Ser Ala Thr Thr Val Lys
Ala Ile Cys Thr Gly Val Ile Asn Met Asn Val Leu Ser Thr Thr Arg
Phe Gln Leu Asn Thr Cys Thr Arg Thr Ser Ile Thr Pro Arg Pro Cys
Pro Tyr Ser Ser Arg Thr Glu Thr Asn Tyr Ile Cys Val Lys Cys Glu
Asn Gln Tyr Pro Val His Phe Ala Gly Ile Gly Arg Cys Pro
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atc atc tgc aac act atc atg gac aac atc tac atc gtt ggt ggt
Ile Ile Cys Asn Thr Ile Met Asp Asn Asn Ile Tyr Ile Val Gly Gly
            20
cag tgc aaa cgt gtt aac act ttc atc atc tct gct act act gtt
Gln Cys Lys Arg Val Asn Thr Phe Ile Ile Ser Ser Ala Thr Thr Val
         35
                             40
aaa gct atc tgc act ggt gtt atc aac atg aac gtt ctg tct act act
Lys Ala Ile Cys Thr Gly Val Ile Asn Met Asn Val Leu Ser Thr Thr
cgt ttc cag ctg aac act tgc act cgt act tct atc act ccg cgt ccg
                                                                  240
Arg Phe Gln Leu Asn Thr Cys Thr Arg Thr Ser Ile Thr Pro Arg Pro
```

By

<210> 24 <211> 110 <212> PRT

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tgc ccg tac tct tct cgt act gaa act aac tac atc tgc gtt aaa tgc
Cys Pro Tyr Ser Ser Arg Thr Glu Thr Asn Tyr Ile Cys Val Lys Cys
gaa aac cag tac ccg gtt cat ttc gct ggt atc ggt cgt tgc ccg
                                                                   333
Glu Asn Gln Tyr Pro Val His Phe Ala Gly Ile Gly Arg Cys Pro
<210> 26
<211> 111
<212> PRT
<213> Rana catesbeiana
<400> 26
Met Ser Asn Trp Ala Thr Phe Gln Gln Lys His Ile Ile Asn Thr Pro
Ile Ile Cys Asn Thr Ile Met Asp Asn Asn Ile Tyr Ile Val Gly Gly
                                 25
             20
Gln Cys Lys Arg Val Asn Thr Phe Ile Ile Ser Ser Ala Thr Thr Val
Lys Ala Ile Cys Thr Gly Val Ile Asn Met Asn Val Leu Ser Thr Thr
Arg Phe Gln Leu Asn Thr Cys Thr Arg Thr Ser Ile Thr Pro Arg Pro
Cys Pro Tyr Ser Ser Arg Thr Glu Thr Asn Tyr Ile Cys Val Lys Cys
Glu Asn Gln Tyr Pro Val His Phe Ala Gly Ile Gly Arg Cys Pro
<210> 27
<211> 2855
<212> DNA
<213> Rana pipiens
<220>
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<222> (97)..(477)
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totottatat ataaaggoot gatoacgact tocaga atg ttt cca aaa tto toa
                                        Met Phe Pro Lys Phe Ser
ttt ctc ctg ata ttt gca gtt gtt ttg agt ctc act cat aag tcc tta
Phe Leu Leu Ile Phe Ala Val Val Leu Ser Leu Thr His Lys Ser Leu
```

15

20

10

B'y

|            |            |            |            |                  |            |            |            |            |                  |            |            |            |            | acc<br>Thr       |            | 210  |
|------------|------------|------------|------------|------------------|------------|------------|------------|------------|------------------|------------|------------|------------|------------|------------------|------------|------|
|            |            |            |            |                  |            |            |            |            |                  |            |            |            |            | tgc<br>Cys       |            | 258  |
|            |            |            |            |                  |            |            |            |            |                  |            |            |            |            | gcc<br>Ala       |            | 306  |
| tgt<br>Cys | aaa<br>Lys | gga<br>Gly | att<br>Ile | ata<br>Ile<br>75 | gcc<br>Ala | tcc<br>Ser | aaa<br>Lys | aat<br>Asn | gtg<br>Val<br>80 | tta<br>Leu | act<br>Thr | acc<br>Thr | tct<br>Ser | gag<br>Glu<br>85 | ttt<br>Phe | 354  |
|            |            |            |            |                  |            |            |            |            |                  |            |            |            |            | aaa<br>Lys       |            | 402  |
|            |            |            |            |                  |            |            |            |            |                  |            |            |            |            | gct<br>Ala       |            | 450  |
|            |            |            |            | ggt<br>Gly       |            |            |            |            | taga             | aaata      | atg t      | ttga       | acaa       | ca               |            | 497  |
| ggga       | atgte      | gat a      | aagca      | agcto            | gc aa      | agaa       | attat      | ttt        | gaag             | gtga       | att        | tacta      | aaa        | gaca             | ctaatt     | 557  |
| ttgo       | cataa      | aat 1      | tttc       | ccas             | ga go      | cttac      | ccggt      | agt        | taaga            | aaaa       | ttc        | caaca      | agg        | gagc             | caagca     | 617  |
| caga       | aaagt      | aa a       | actaa      | aggag            | ge ea      | aagt       | aatt       | ata        | aaaa             | gtca       | cact       | tgga       | ccg        | ctgc             | tactgc     | 677  |
| acto       | cagat      | ga (       | ccaaa      | atgag            | ga aa      | acaga      | acaaa      | a aad      | cagca            | agag       | ttgg       | ggaag      | gcg        | cagat            | ccggg      | 737  |
| aggt       | ggcg       | 199 (      | gagto      | caatt            | g gg       | ggate      | ggagt      | c cca      | atgte            | gaga       | tttg       | ggaa       | ccg        | tttgi            | ttgctg     | 797  |
| gtga       | aagca      | atg 1      | tggc       | eggte            | gc ac      | cagta      | acaca      | a tgg      | gggaa            | aaga       | tagi       | tcgga      | att :      | ggcc             | gggctc     | 857  |
| gctg       | gtggt      | gg 1       | tgcc       | ggcgg            | gt to      | gagco      | caaag      | ggt        | ggtgg            | ggga       | gate       | ggct       | gtc        | cccc             | cttctg     | 917  |
| tggg       | gggct      | gt (       | ggaca      | agagg            | gg ag      | gctgo      | cggad      | cag        | ggggt            | tggg       | aggo       | cctg       | gag        | agaat            | tttca      | 977  |
| aaca       | agcto      | gac q      | gtgg       | ccggg            | gg ct      | ggg        | cagca      | a tog      | 3999             | aggg       | gaag       | gggct      | gg         | gctca            | agatcc     | 1037 |
| agga       | agca       | atg (      | gtcad      | ctgta            | at ga      | accaç      | gagto      | g gaa      | agat             | ggca       | gago       | ccgct      | .gc        | agtg             | gccggg     | 1097 |
| gaga       | accag      | gag g      | ggato      | ctgtg            | gc co      | cagco      | ctttc      | ccc        | ctcc             | ctga       | tgt        | ggcc       | egt        | ttttg            | ggttat     | 1157 |
| ggta       | acco       | gct (      | ccca       | gctgt            | t to       | 99999      | gtgtt      | tt.        | eggg             | cttc       | gcat       | tttt       | gg         | tctg             | eggete     | 1217 |
| ccto       | ctgto      | cca (      | cggc       | cctca            | at gg      | gagg       | 39999      | g tgg      | ggcat            | tttc       | tcca       | accgo      | cct        | ttgg             | ctctgt     | 1277 |
| tgct       | ggca       | act 9      | gtcg       | cagco            | ga gt      | ttg        | gccag      | g tca      | atggo            | ctca       | ttti       | tccca      | att        | tgtca            | atgtgt     | 1337 |
| gtts       | gtt        | gca 1      | tgtt       | ttgto            | g go       | ggtg       | ggact      | gti        | tttga            | aatt       | tcad       | catg       | gat        | tcca             | tcttcg     | 1397 |

gttggttcct tgccacctcc tggatctgtg ctttccaatt ctgttttttc cccagcgctt 1457 agtggatgca gtgaaactct ggtgattacc atcatccaat catgtgcaag aaaaaatatt 1517 ttcatatttc ttccacccaa ttgggtattc attaggaagt ttgagcacat tcacgttcta 1577 gggaaaatga gtgcaactgc acttccaaag ttcacagtct atttgccttt agtaaatcca 1637 ccccattatt tctgagcaga ggacaaatct atggcaacaa aaaaacttta cctactgaat 1697 tattttatat tgattgaaga taatetttet tteattteet aaatattgta ateaaaatta 1757 atacataaca gctatgtatt ataccacagc agcaaatgtt aaaatagttt taaacgtaaa 1817 atatgtttta cettaaagtg gaagtaaact tetateaeta aattttaeet ataggtgaga 1877 cccatgcgct cttcaggaat ggccgctggt gctgttcctt cagagccctg tgctgcgaac 1937 ggcggctccc gtgtgcatgt acaggagtga cgtcatcaca gctccggcca gtcacagagt 1997 tagagttcaa gtgtgagtgg cttgagccac gatgatgtcg ctcccaaaca tgtgtgcggg 2057 ggtctccgtt tgcggcgcag gacactgggg gaatagcatg ggtgtgccgt tccttcagag 2117 catatgcgtg ggtgacgtca ctagctgcat ctaaagtaat atctcctaaa caatgcacat 2177 ttaggagata gttacagtac ctatgggtaa gccttattgt aggcttacct ataggtaaaa 2237 atcatgcatg ggagtttact tccactttaa caccggcata gctgcgtgtt tataaaccaa 2297 ccagcagtag caaatgtagg gatgaggaga gcaggctgac atattaaagt aaaaatctta 2357 cctatagtgg ttgaaagtag ttgaaaataa gatggcctgc agggtcttaa aaaggctagg 2417 atagcacagt atccacatga ggcaccagat ctcgctcccc cacacatgag tagcaaggag 2477 caatggtaat gtgagtttet taggetegae egttaaatag egttggeeet eeaagtgata 2537 catgggagat aagcagatgt ccgcgtatgc acgcagacat atgtgggcgg atgttgggat 2597 aggacgatca gagagatgct cagatctgcc cgaaggagaa aggtggaaac atccattcaa 2657 tgtcatatgc ctaaagaagc cacccaccat aaaaagttaa tagatcatca ggtggcagcc 2717 aaccacaca ggcccaaagg agggtggccc cagtggaccg tataggaaca gcactcagct 2777 atcacataat tacacaagag tatagagacc cattgtgggt attaacaacc aaatggctaa 2837 aaaaaaaaa aaaaaaaa 2855



aaaaaaa

<sup>&</sup>lt;210> 28 <211> 127

<sup>&</sup>lt;212> PRT

<sup>&</sup>lt;213> Rana pipiens

Met Phe Pro Lys Phe Ser Phe Leu Leu Ile Phe Ala Val Val Leu Ser

<400> 28

<213> Artificial Sequence

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Leu Thr His Lys Ser Leu Cys Gln Asp Trp Leu Thr Phe Gln Lys Lys
His Leu Thr Asn Thr Arg Asp Val Asp Cys Asn Asn Ile Met Ser Thr
Asn Leu Phe His Cys Lys Asp Lys Asn Thr Phe Ile Tyr Ser Arg Pro
Glu Pro Val Lys Ala Ile Cys Lys Gly Ile Ile Ala Ser Lys Asn Val
Leu Thr Thr Ser Glu Phe Tyr Leu Ser Asp Cys Asn Val Thr Ser Arg
Pro Cys Lys Tyr Lys Leu Lys Lys Ser Thr Asn Thr Phe Cys Val Thr
            100
                                105
                                                     110
Cys Glu Asn Gln Ala Pro Val His Phe Val Gly Val Gly His Cys
                            120
<210> 29
<211> 4
<212> PRT
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<400> 29
Cys Val Ile Met
<210> 30
<211> 27
<212> DNA
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<220>
<223> Description of Artificial Sequence: Primer
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<210> 31
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<212> DNA
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<220>
<223> Description of Artificial Sequence: Primer
<400> 31
                                                                   27
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<210> 32
<211> 96
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Synthetic
      oligonucleotide
<400> 32
cagaactggg ctactttcca gcagaaacat atcatcaaca ctccgatcat ctgcaacact 60
atcatggaca acaacatcta catcgttggt ggtcag
                                                                   96
<210> 33
<211> 86
<212> DNA
<213> Artificial Sequence
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<223> Description of Artificial Sequence: Synthetic
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<400> 33
tacategttg gtggtcagtg caaacgtgtt aacactttca teatetetet getactaetg 60
ttaaacgtat ctgcactggt gttatc
<210> 34
<211> 96
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Synthetic
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<400> 34
atctgcactg gtgttactaa catgaacgtt ctgtctacta ctcgtttcca gctgaacact 60
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<210> 35
<211> 21
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Synthetic
      oligonucleotide
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But

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<400> 35
                                                                   21
gttgataaca ccagtgcaga t
<210> 36
<211> 21
<212> DNA
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<400> 36
                                                                   21
atctgcactg gtgttatcaa c
<210> 37
<211> 95
<212> DNA
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<400> 37
actecgegte egtgeeegta etettetegt actgaaacta actacatetg egttaaatge 60
gaaaaccagt acccggttca tttcgctggt atcgg
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<212> DNA
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ctactttcca g
<210> 39
<211> 48
<212> DNA
<213> Artificial Sequence
<220>
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<400> 39
cgcgccggat ccctactacg ggcaacgacc gataccagcg aaatgaac
                                                                   48
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<210> 40
<211> 96
<212> DNA
<213> Artificial Sequence
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atcctgcaga acaacatcta catcgttggt ggtcag
<210> 41
<211> 96
<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence: Synthetic
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<400> 41
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tgcactcgta cttctatcac tccgcgtccg tgcccg
<210> 42
<211> 33
<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence: Primer
<400> 42
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ggattccata tgcagaactg ggctattttc cag
<210> 43
<211> 6
<212> PRT
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<220>
<223> Description of Artificial Sequence: Synthetic
      6X His tag
<400> 43
His His His His His
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BI